

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference	FOR FURTHER see Notification o	f Transmittal of International Search Report 20) as well as, where applicable, item 5 below.
	ACTION (Form PC1/ISA/2	zoj as well as, miste appresant, nem se
038602/0272 International application No.	International filing date (day/month/year)	(Earliest) Priority Date (day/month/year)
	30/08/2000	30/08/1999
PCT/US 00/23744	30/06/2000	20,00,200
Applicant		
NEW YORK UNIVERSITY SCHOOL	OF MEDICINE et al.	
NEW YURK UNIVERSITY SCHOOL	Of FILDIOTICE CO WIT	
This International Search Report has been according to Article 18. A copy is being tra	n prepared by this International Searching Auth Insmitted to the International Bureau.	nority and is transmitted to the applicant
This International Search Report consists  It is also accompanied by	of a total of sheets. a copy of each prior art document cited in this	report.
Basis of the report		
With regard to the language, the language in which it was filed, unline	international search was carried out on the baless otherwise indicated under this item.	
Authority (Rule 23 1(b)).	vas carried out on the basis of a translation of t	
b. With regard to any nucleotide an was carried out on the basis of the	nd/or amino acid sequence disclosed in the in e sequence listing:	nternational application, the international search
contained in the internation	onal application in written form.	•
filed together with the inte	ernational application in computer readable for	m.
	o this Authority in written form.	
furnished subsequently to	o this Authority in computer readble form.	
the statement that the su	bsequently furnished written sequence listing of as filed has been furnished.	
the statement that the inf	formation recorded in computer readable form	is identical to the written sequence listing has been
2. X Certain claims were for	und unsearchable (See Box I).	
3. X Unity of invention is la		
4. With regard to the title,	when the desired by the englisher	
the text is approved as s	submitted by the applicant.	
the text has been establ	ished by this Authority to read as follows:	
5. With regard to the abstract,	·	
TVT the tout is approved as s	submitted by the applicant.	
	ished, according to Rule 38.2(b), by this Autho he date of mailing of this international search ro	ority as it appears in Box III. The applicant may, eport, submit comments to this Authority.
	blished with the abstract is Figure No.	
as suggested by the ap		X None of the figures.
because the applicant fa	ailed to suggest a figure.	
	er characterizes the invention.	

Form PCT/ISA/210 (first sheet) (July 1998)

International application No. PCT/US 00/23744

B x I Obs rvations where c rtain claims were found unsearchable (Continuation of item 1 of first sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:  see FURTHER INFORMATION sheet PCT/ISA/210
2. X Claims Nos.: 66 because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:  See FURTHER INFORMATION sheet PCT/ISA/210
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
B x II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
see additional sheet
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:  Claims 1-27, 43-51 and 54-86.
Remark on Protest  The additional search fees were accompanied by the applicant's protest.  No protest accompanied the payment of additional search fees.

#### FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.1

Although claim 68 is directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.

Although claims 43-51 could be considered as a mere presentation of information or a computerprogram, Rule 39.1(v and vi) PCT, the search has been carried out as far as pssible in our systematic documentation

Continuation of Box I.2

Claims Nos.: 66

Present claim 66 relates to a product defined by reference to a desirable characteristic or property, namely modulation of receptor protein tyrosine kinase.

The claims cover all products having this characteristic or property, whereas the application provides support within the meaning of Article 6 PCT and/or disclosure within the meaning of Article 5 PCT for no such product. In the present case, the claims so lack support, and the application so lacks disclosure, that a meaningful search over the whole of the claimed scope is impossible. Independent of the above reasoning, the claims also lack clarity (Article 6 PCT). An attempt is made to define the product/compound/method/apparatus by reference to a result to be achieved. Again, this lack of clarity in the present case is such as to render a meaningful search over the whole of the claimed scope impossible. Consequently, the search has not been carried out.

The applicant's attention is drawn to the fact that claims, or parts of claims, relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure.

#### FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. Claims: 1-27 43-51 54-86

2. Claims: 28-42 52 53 87-89

Crystals comprising a receptor binding core of a stem cell factor and there use

International Application No PAT/US 00/23744

A. CLASSIFICATION OF SUBJECT MATI G01N33/68

G06F17/30

C12Q1/37

A61K38/00

According to International Patent Classification (IPC) or to both national classification and IPC

#### **B. FIELDS SEARCHED**

 $\begin{array}{ll} \mbox{Minimum documentation searched} & \mbox{(classification system followed by classification symbols)} \\ IPC 7 & C12N & C07K & A61K & C12Q & G01N \end{array}$ 

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, BIOSIS, WPI Data

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	PELLEGRINI L ET AL: "The role of heparin in the complex formation between fibroblast growth factor 2 and its high affinity receptor: Comparative modelling and biochemical studies." BIOCHEMICAL SOCIETY TRANSACTIONS, vol. 26, no. 3, August 1998 (1998-08), pages 545-549, XP000993500 665th Meeting of the Biochemical Society; Southampton, England, UK; March 31-April 2, 1998 ISSN: 0300-5127 the whole document	1-27, 43-51, 54-65, 67-86
V Furth	ner documents are listed in the continuation of box C.	are listed in annex.

X Further documents are listed in the continuation of box C.	X Patent family members are listed in annex.
<ul> <li>Special categories of cited documents:</li> <li>"A" document defining the general state of the art which is not considered to be of particular relevance</li> <li>"E" earlier document but published on or after the international filling date</li> <li>"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)</li> <li>"O" document referring to an oral disclosure, use, exhibition or other means</li> <li>"P" document published prior to the international filing date but later than the priority date claimed</li> </ul>	"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention  "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone  "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.  "8" document member of the same patent family
Date of the actual completion of the international search	Date of mailing of the international search report
26 July 2001	2 6. 10. 01
Name and mailing address of the ISA  European Patent Office, P.B. 5818 Patentlaan 2  NL - 2280 HV Rijswijk  Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,  Fax: (+31-70) 340-3016	Authorized officer  VAN DER SCHAAL C.A.

Form PCT/ISA/210 (second sheet) (July 1992)

International Application No PCI/US 00/23744

C.(Continu	ation) DOCUMENTS CONSIDE	
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Υ	HUHTALA MIKKO T ET AL: "A dimeric ternary complex of FGFR1, heparin and FGF-1 leads to an 'electrostatic sandwich' model for heparin binding." STRUCTURE (LONDON), vol. 7, no. 6, June 1999 (1999-06), pages 699-709, XP000993253 ISSN: 0969-2126 the whole document	1-27, 43-51, 54-65, 67-86
Υ	WO 98 07835 A (SCHLESSINGER JOSEPH ;LIANG CONGXIN (US); SUGEN INC (US); TANG PENG) 26 February 1998 (1998-02-26) cited in the application the whole document	1-27, 43-51, 54-65, 67-86
Y	GIVOL DAVID ET AL: "Complexity of FGF receptors: Genetic basis for structural diversity and functional specificity." FASEB (FEDERATION OF AMERICAN SOCIETIES FOR EXPERIMENTAL BIOLOGY), vol. 6, no. 15, 1992, pages 3362-3369, XP002167375 1992 ISSN: 0892-6638 cited in the application the whole document	1-27, 43-51, 54-65, 67-86
Υ	LIEKENS SANDRA ET AL: "Modulation of fibroblast growth factor-2 receptor binding, signaling, and mitogenic activity by heparin-mimicking polysulfonated compounds."  MOLECULAR PHARMACOLOGY, vol. 56, no. 1, July 1999 (1999-07), pages 204-213, XP000993567 ISSN: 0026-895X the whole document	58-65
P,X	PLOTNIKOV ALEXANDER N ET AL: "Structural basis for FGF receptor dimerization and activation." CELL, vol. 98, no. 5, 3 September 1999 (1999-09-03), pages 641-650, XP002167376 ISSN: 0092-8674	1-27, 43-51, 54-57, 67-86
Υ	the whole document/	58-65

International Application No PCT/US 00/23744

P,X PLOTNIKOV ALEXANDER N ET AL: "Crystal structures of two FGF-FGFR complexes 43-51, reveal the determinants of ligand-receptor 54-57, specificity." 67-69, CELL, vol. 101, no. 4, 12 May 2000 (2000-05-12), pages 413-424, XP002167377 ISSN: 0092-8674 the whole document 58-65, 70-75  P,X PELLEGRINI LUCA ET AL: "Crystal structure of fibroblast growth factor receptor ectodomain bound to ligand and heparin." 43-51, NATURE (LONDON), vol. 407, no. 6807, 2000, pages 1029-1034, XP002167378 ISSN: 0028-0836 the whole document 5-7, 9-18, 24-27, 54-65, 75,77-86  P,X STAUBER DEBORAH J ET AL: "Structural interactions of fibroblast growth factor factor receptor with its ligands." 43-51, PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES OF THE UNITED STATES, vol. 97, no. 1, 4 January 2000 (2000-01-04), pages 49-54, XP002167379 Jan. 4, 2000 ISSN: 0027-8424		ation) DOCUMENTS CONSIDE O BE RELEVANT	7
Structures of two FGF-FGFR complexes reveal the determinants of ligand-receptor specificity."	Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y the whole document 58-65, 70-75  P,X PELLEGRINI LUCA ET AL: "Crystal structure of fibroblast growth factor receptor ectodomain bound to ligand and heparin." 43-51, NATURE (LONDON), vol. 407, no. 6807, 2000, pages 1029-1034, XP002167378	P,X	structures of two FGF-FGFR complexes reveal the determinants of ligand-receptor specificity." CELL, vol. 101, no. 4, 12 May 2000 (2000-05-12), pages 413-424, XP002167377	43-51, 54-57, 67-69,
of fibroblast growth factor receptor ectodomain bound to ligand and heparin." NATURE (LONDON), vol. 407, no. 6807, 2000, pages 1029-1034, XP002167378 ISSN: 0028-0836 the whole document  P,X  STAUBER DEBORAH J ET AL: "Structural interactions of fibroblast growth factor receptor with its ligands." PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES OF THE UNITED STATES, vol. 97, no. 1, 4 January 2000 (2000-01-04), pages 49-54, XP002167379 Jan. 4, 2000 ISSN: 0027-8424 the whole document  19-23, 43-51, 67-74,76 67-74,76 67-74,76 5-7, 9-18, 24-27, 54-65,	Υ		
Y the whole document  5-7, 9-18, 24-27, 54-65, 75,77-86  P,X STAUBER DEBORAH J ET AL: "Structural interactions of fibroblast growth factor receptor with its ligands." PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES OF THE UNITED STATES, vol. 97, no. 1, 4 January 2000 (2000-01-04), pages 49-54, XP002167379 Jan. 4, 2000 ISSN: 0027-8424 Y the whole document  5-7, 9-18, 24-27, 54-65,	P,X	of fibroblast growth factor receptor ectodomain bound to ligand and heparin." NATURE (LONDON), vol. 407, no. 6807, 2000, pages 1029-1034, XP002167378	19-23, 43-51,
interactions of fibroblast growth factor receptor with its ligands." PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES OF THE UNITED STATES, vol. 97, no. 1, 4 January 2000 (2000-01-04), pages 49-54, XP002167379 Jan. 4, 2000 ISSN: 0027-8424 the whole document  5-7, 9-18, 24-27, 54-65,	Y		9-18, 24-27, 54-65,
Y the whole document 5-7, 9-18, 24-27, 54-65,	P,X	interactions of fibroblast growth factor receptor with its ligands." PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES OF THE UNITED STATES, vol. 97, no. 1, 4 January 2000 (2000-01-04), pages 49-54, XP002167379 Jan. 4, 2000	19-23, 43-51,
	Y		9-18, 24-27, 54-65,
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Information on patent family members

International Application No

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
WO 9807835 A	26-02-1998	US 5942428 A AU 733890 B AU 4160397 A EP 0931152 A JP 2001514484 T	24-08-1999 31-05-2001 06-03-1998 28-07-1999 11-09-2001